

<120> Use of Labelled CCK-B Receptor Ligands for the
Detection and Localization of Malignant Human Tumours

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Detection and Localization of Malignant Human Tumours
<130> 1668-303
<140>09/125,823
<141> 1999-01-19
<150> EPO 96200498.2
<151> 1996-02-27
<150> PCT/US97/03056
<151> 1997-02-25
<160>27
<170> PatentIn Ver. 2.0
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<223> The peptide is labelled with a radionuclide or
      with a paramagnetic metal isotope.
<220>
<221> MOD RES
<222> (7)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400>1
Tyr Met Gly Trp Met Asp Xaa
<210>2
<211>8
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<223> The peptide is labelled with a radionuclide or with a paramagnetic metal isotope.

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<220>
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<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
       analog.
<400>2
Asp Tyr Met Gly Trp Met Asp Xaa
<210>3
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<221> MOD_RES
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      with a paramagnetic metal isotope.
<220>
<221> MOD_RES
<222> (3)
<223> Nle
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<221> MOD RES
<222>(6)
<223> Nle
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<221> MOD RES
<222> (8)
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<400> 3
Asp Tyr Xaa Gly Trp Xaa Asp Xaa
                 5
<210>4
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<221> MOD_RES
<222>(1)..(8)
<223> The peptide is labelled with a radionuclide or
      with a paramagnetic metal isotope.
<220>
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<221> MOD RES
<222>(1)
<223> Xaa is DAsp.
<220>
<221> MOD RES
<222>(3)
<223> Nle
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<221> MOD_RES
<222>(6)
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<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400>4
Xaa Tyr Xaa Gly Trp Xaa Asp Xaa
<210>5
<211>8
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<223> The peptide is labelled with a radionuclide or
      with a paramagnetic metal isotope.
<220>
<221> MOD_RES
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<223> Xaa is DAsp.
<220>
<221> MOD_RES
<222> (8)
<223> Xaa is Phe-NH2.
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
Xaa Tyr Met Gly Trp Met Asp Xaa
  1
<210>6
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<211>8

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<212> PRT
<213> Artificial Sequence
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<221> MOD RES
<222>(1)..(8)
<223> The peptide is labelled with a radionuclide or
      with a paramagnetic metal isotope.
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<221> MOD_RES
<222>(3)
<223> Nle
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<222> (6)
<223> Nle
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<221> MOD RES
<222> (8)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400>6
Xaa Tyr Xaa Gly Trp Xaa Asp Xaa
<210>7
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      with a paramagnetic metal isotope.
<220>
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<223> Nle
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<222> (8)
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<223> Xaa is Phe-NH2.
<220>
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      analog.
<400> 7
Asp Tyr Thr Gly Trp Xaa Asp Xaa
<210>8
<211>9
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<213> Artificial Sequence
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<221> MOD_RES
<222>(1)..(9)
<223> The peptide is labelled with a radionuclide or
      with a paramagnetic metal isotope.
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<222> (4)
<223> Nle
<220>
<221> MOD_RES
<222>(7)
<223> Nle
<220>
<221> MOD_RES
<222> (9)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400>8
Arg Asp Tyr Xaa Gly Trp Xaa Asp Xaa
                  5
  1
<210>9
<211>9
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD RES
<222>(1)..(9)
<223> The peptide is labelled with a radionuclide or
       with a paramagnetic metal isotope.
<220>
<221> MOD RES
<222>(7)
<223> Nle
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<220>
<221> MOD_RES
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<223> Xaa is Phe-NH2.
<220>
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<400>9
Arg Asp Tyr Thr Gly Trp Xaa Asp Xaa
<210> 10
<211>10
<212> PRT
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<221> MOD_RES
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      with a paramagnetic metal isotope.
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<223> Nle
<220>
<221> MOD_RES
<222>(8)
<223> Nle
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<221> MOD_RES
<222>(10)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
     analog.
<400> 10
Tyr Gly Asp Tyr Xaa Gly Trp Xaa Asp Xaa
                 5
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<210>11
<211>10
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222>(1)..(10)
<223> The peptide is labelled with a radionuclide or
      with a paramagnetic metal isotope.
<220>
<221> MOD_RES
<222>(1)
<223> Xaa is DTyr.
<220>
<221> MOD_RES
<222>(5)
<223> Nle
<220>
<221> MOD_RES
<222>(8)
<223> Nle
<220>
<221> MOD_RES
<222> (10)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
       analog.
<400> 11
Xaa Gly Asp Tyr Xaa Gly Trp Xaa Asp Xaa
   1
<210>12
<211>10
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222>(1)
<223> Xaa is DTyr.
 <220>
 <221> MOD_RES
 <222>(5)
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 <222>(8)
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<220>
<221> MOD_RES
<222> (10)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 12
Xaa Gly Asp Tyr Xaa Gly Trp Xaa Asp Xaa
                 5
<210>13
<211>10
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222> (1)
<223> Xaa is 125I iodinated DTyr.
<220>
<221> MOD RES
<222>(5)
<223> Nle
<220>
<221> MOD_RES
<222> (8)
<223> Nle
<220>
<221> MOD_RES
<222>(10)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 13
Xaa Gly Asp Tyr Xaa Gly Trp Xaa Asp Xaa
                  5
  1
<210> 14
<211>10
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222> (1)
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<223> Xaa is DTyr.

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<220>
<221> MOD_RES
<222> (4)
<223> Xaa is 125I iodinated Tyr.
<220>
<221> MOD_RES
<222> (5)
<223> Nle
<220>
<221> MOD_RES
<222> (8)
<223> N1e
<220>
<221> MOD_RES
<222>(10)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 14
Xaa Gly Asp Xaa Xaa Gly Trp Xaa Asp Xaa
  1
                 5
<210>15
<211>10
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222> (1)
<223> Xaa is 125I iodinated DTyr.
<220>
<221> MOD_RES
<222> (4)
<223> Xaa is Tyr(SO3H).
<220>
<221> MOD_RES
<222> (5)
<223> Nle
<220>
<221> MOD_RES
<222>(8)
<223> Nle
<220>
<221> MOD RES
<222>(10)
<223> Xaa is Phe-NH2.
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<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 15
Xaa Gly Asp Xaa Xaa Gly Trp Xaa Asp Xaa
  1
                  5
<210> 16
<211>8
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD RES
<222> (2)
<223> Xaa is Tyr(SO3H).
<220>
<221> MOD_RES
<222> (8)
<223> Xaa is Phe-NH2.
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 16
Asp Xaa Met Gly Trp Met Asp Xaa
<210>17
<211>8
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD RES
<222> (8)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 17
Asp Tyr Met Gly Trp Met Asp Xaa
                 5
 1
<210>18
<211>10
<212> PRT
<213> Artificial Sequence
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<220>
<221> MOD_RES
<222>(1)
<223> Xaa is DTyr.
<220>
<221> MOD RES
<222> (4)
<223> Xaa is Tyr(SO3H).
<220>
<221> MOD_RES
<222>(5)
<223> Nle
<220>
<221> MOD_RES
<222>(8)
<223> Nle
<220>
<221> MOD RES
<222> (10)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 18
Xaa Gly Asp Xaa Xaa Gly Trp Xaa Asp Xaa
                  5
<210>19
<211>8
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222>(1)
<223> Xaa is DTPA substituted Asp.
<220>
<221> MOD RES
<222>(8)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 19
Xaa Tyr Met Gly Trp Met Asp Xaa
  1
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<210> 20
<211>8
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222>(1)
<223> Xaa is DTPA substituted Asp.
<220>
<221> MOD_RES
<222>(3)
<223> Nle
<220>
<221> MOD_RES
<222>(6)
<223> Nle
<220>
<221> MOD_RES
<222> (8)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400>20
Xaa Tyr Xaa Gly Trp Xaa Asp Xaa
<210>21
<211>8
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222> (1)
<223> Xaa is DTPA substituted DAsp.
<220>
<221> MOD_RES
<222>(3)
<223> Nle
<220>
<221> MOD_RES
<222>(6)
<223> Nle
<220>
<221> MOD_RES
<222> (8)
<223> Xaa is Phe-NH2.
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<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 21
Xaa Tyr Xaa Gly Trp Xaa Asp Xaa
                 5
  1
<210>22
<211>8
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD RES
<222>(1)
<223> Xaa is DTPA substituted DAsp.
<220>
<221> MOD_RES
<222> (8)
<223> Xaa is Phe-NH2.
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 22
Xaa Tyr Met Gly Trp Met Asp Xaa
  1
                 5
<210> 23
<211>8
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD RES
<222> (1)
<223> Xaa is beta-DTPA substituted Dpr.
<220>
<221> MOD_RES
<222>(3)
<223> Nle
<220>
<221> MOD_RES
<222>(6)
<223> Nle
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<221> MOD_RES
<222> (8)
<223> Xaa is Phe-NH2.
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<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 23
Xaa Tyr Xaa Gly Trp Xaa Asp Xaa
<210> 24
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<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222>(1)
<223> Xaa is DTPA substituted Asp.
<220>
<221> MOD_RES
<222> (6)
<223> Nle
<220>
<221> MOD RES
<222> (8)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
     analog.
<400> 24
Xaa Tyr Thr Gly Trp Xaa Asp Xaa
  1
<210> 25
<211>8
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD RES
<222> (1)
<223> Xaa is ll5Indium-DTPA substituted Asp.
<220>
<221> MOD_RES
<222>(3)
<223> Nle
<220>
<221> MOD_RES
<222>(6)
<223> Nle
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<220>
<221> MOD_RES
<222> (8)
<223> Xaa is Phe-NH2.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 25
Xaa Tyr Xaa Gly Trp Xaa Asp Xaa
<210> 26
<211>8
<212> PRT
<213> Artificial Sequence
<220>
<221> MOD_RES
<222> (1)
<223> Xaa is ll5Indium-DTPA substituted DAsp.
<220>
<221> MOD_RES
<222>(3)
<223> Nle
<220>
<221> MOD_RES
<222>(6)
<223> Nle
<220>
<221> MOD_RES
<222> (8)
<223> Xaa is Phe-NH2.
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 26
Xaa Tyr Xaa Gly Trp Xaa Asp Xaa
                  5
  1
<210>27
<211>33
<212> PRT
<213> Artificial Sequence
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<220>
<221> SITE
<222>(1)..(25)
<223> The first 25 Xaa's may or may not be present; can
      be equal or different; are selected from Ala, Leu,
      Asn, Dpr, Gln, Glu, Ser, Ile, Met, His, Asp, Lys,
      Gly, Thr, Pro, Pyr, Arg, Tyr, Trp, Val and Phe.
<220>
<221> SITE
<222> (26)
<223> Xaa is Asp, Dpr, Glu or Pyr, with the proviso that
      Xaa can only be Pyr when residues 1-25 are not
      present.
<220>
<221> SITE
<222> (28)
<223> Xaa is Met, Leu or Nle.
<220>
<221> SITE
<222> (31)
<223> Xaa is Met, Leu or Nle.
<220>
<221> SITE
<222> (33)
<223> Xaa is Phe terminating in a hydroxy group, an
      acetoxy group or an amino group.
<220>
<223> Description of Artificial Sequence: Cholecystokinin
      analog.
<400> 27
Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Xaa Tyr Xaa Gly Trp Xaa Asp
                                25
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Xaa